# (19) World Intellectual Property Organization

International Bureau





(43) International Publication Date 8 July 2004 (08.07.2004)

PCT

## (10) International Publication Number WO 2004/056982 A3

(51) International Patent Classification<sup>7</sup>: C12N 9/12, 5/10, G01N 33/53, C07K 16/40

(21) International Application Number:

PCT/EP2003/014674

(22) International Filing Date:

19 December 2003 (19.12.2003)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

0230014.3 60/436,380 23 December 2002 (23.12.2002) GB

0230014.3

23 December 2002 (23.12.2002) US

- (71) Applicant (for all designated States except US): DEVGEN NV [BE/BE]; Technologiepark 30, B-9052 Gent-Zwij-naarde (BE).
- (72) Inventor; and
- (75) Inventor/Applicant (for US only): DE WILDE, Gert, Jules, Hector [BE/BE]; Dr. Armand Rubbensstraat 25, B-9240 Zele (BE).
- (74) Agents: MURPHY, Colin, Damien et al., Boult Wade Tennant, 70 Gray's Inn Road, London WC1X 8BT (GB).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU,

CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (regional): ARIPO patent (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

#### Declaration under Rule 4.17:

- of inventorship (Rule 4.17(iv)) for US only

#### Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- (88) Date of publication of the international search report:
  16 September 2004

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: KINASE SEQUENCES

(57) Abstract: The present invention relates to nucleotide sequences that encode and may be used to express amino acid sequences that are useful in the identification and development of compounds with activity as pharmaceuticals, in particular of compounds for the prevention and treatment of metabolic diseases such as diabetes and obesity. The invention also relates to the amino acid sequences - such as proteins and polypeptides - that are encoded by, and that may be obtained by suitable expression of, the nucleotide sequences of the invention, particularly the amino acid sequences of J1K, PSK, TAO1 and Q9P2I6. The invention also relates to various uses and modulators of, and methods incorporating, the nucleotide and amino acid sequences of the invention.

2004/056982 A3

International Application No FET/EP .03/14674

A. CLASSIFICATION OF SUBJECT MATTER
IPC. 7 C12N9/12 C12N5/10

G01N33/53

C07K16/40

According to International Patent Classification (IPC) or to both national classification and IPC

Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

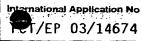
EPO-Internal, BIOSIS, EMBL, GENSEQ, SEQUENCE SEARCH, PAJ, WPI Data, CHEM ABS Data

ategory °	Citation of document, with indication, where appropriate, of the relevant passages  Relevant to claim No.
	DATABASE EMBL 'Online! 1-10.17
	16 September 1995 (1995-09-16)
	WILSON R. ET AL.: "Caenorhabditis elegans
	putative serine/threonine protein kinase
	(Sulu) mRNA, complete cds."
	Database accession no. U32275
	XP002285648
	the whole document
	the whole document
X	DATABASE UNIPROT 'Online! 1-10,17
^	1 November 1995 (1995-11-01)
	COPE M.J.T.V., KENDRICK-JONES A.: "
	Serine/threonine-protein kinase SULU (EC
	2.7.1.37)."
	Database accession no. P46549
	XP002285649
	the whole document
	the whole document
	_/
	the control of the co

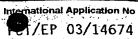
X Further documents are listed in the con	linuation of box C.	Patent family members are listed in annex.
Special categories of cited documents:  'A' document defining the general state of the considered to be of particular relevance  'E' earlier document but published on or after filing date  'L' document which may throw doubts on prio which is cited to establish the publication citation or other special reason (as special document referring to an oral disclosure, to other means  'P' document published prior to the internation later than the priority date claimed	the international  ity claim(s) or date of another lied)  se, exhibition or little filling date but	T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention.  X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone.  Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combined with one or more other such documents, such combined with one or more other such documents, such combined being obvious to a person skilled in the art.
Date of the actual completion of the internation 25 June 2004	al search	Date of mailing of the international search report $12/07/2004$
Name and mailing address of the ISA  European Patent Office, P.B.  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 3  Fax: (+31-70) 340-3016		Authorized officer  Morawetz, R

International Application No T/EP 03/14674

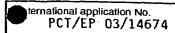
:.(Continu'	ation) DOCUMENTS CONSIDERED TO BE RELEVANT			4 1 4 4 4
<u> </u>			In the second	
category °	Citation of document, with indication, where appropriate, of the relevant passages	, <del>1</del> .	Relevant to claim No.	0
,	DEDMAN V C ET Al. This 10 2 C alors		1.10.17	
(	BERMAN K S ET AL: "kin-18, a C. elegans protein kinase involved in feeding"		1-10,17	
	GENE: AN INTERNATIONAL JOURNAL ON GENES			•
	AND GENOMES, ELSEVIER SCIENCE PUBLISHERS,		}	
	BARKING, GB,	•	,	
	vol. 279, no. 2,			
•	28 November 2001 (2001-11-28), pages		<b>.</b>	
	137-147, XP004329111			
	ISSN: 0378-1119 DN VIC FILE		1.	
	cited in the application			: -
	the whole document			
•				
(	WO 99/53036 A (SUGEN INC ;WHYTE DAVID		1-10,17	
	(US); PLOWMAN GREGORY (US); MARTINEZ		]	
	RICARD) 21 October 1999 (1999-10-21)			
	the whole document	: .		
. 1	4.2 0.0 1.00	•		
(	DATABASE EMBL 'Online!		1-10,17	
	19 January 2000 (2000-01-19)			٠,
	CARTER T.G. ET AL.: "Homo sapiens			
	serine/threonine kinase (KDS) mRNA,		ļ	
	complete cds."			
	Database accession no. AF181985			
ı	XP002285650			
	the whole document			
	the whole document	· · ·	:	
(	TASSI E ET AL: "Human JIK, a novel member		1-10,17	
`	of the STE20 kinase family that inhibits		1 10,17	
	JNK and is negatively regulated by			
	epidermal growth factor"			•
		D.		٠.
	JOURNAL OF BIOLOGICAL CHEMISTRY, AMERICAN	•		
	SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE,			
	MD, US,		; · · ·	
	vol. 274, no. 47,	•		
	19 November 1999 (1999-11-19), pages		l ·	
	33287-33295, XP002965003			
	ISSN: 0021-9258		1	
	cited in the application			
	the whole document		· ·	
,	7UANC METRING ET AL. "Closing of DDV o		1 10 17	
(	ZHANG WEIPING ET AL: "Cloning of DPK, a		1-10,17	
į	Minovel dendritic cell-derived protein		1	
	kinase activating the ERK1/ERK2 and			
	JNK/SAPK pathways"	,	į	
	BIOCHEMICAL AND BIOPHYSICAL RESEARCH		1	.5
	COMMUNICATIONS,		l	
	vol. 274, no. 3,		j 4.	
	11 August 2000 (2000-08-11), pages		I	
	872-879, XP002285645		} ·	
	ISSN: 0006-291X		· .	
	cited in the application		1	
	the whole document			
			<b>.</b>	
	-/		İ	
		•	ļ.	
1			i .	



C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	FEI/EP U.	3, 140, 4
Cat y	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
* #			
X	MOORE TANYA M ET AL: "PSK, a novel STE20-like kinase derived from prostatic carcinoma that activates the c-Jun N-terminal kinase mitogen-activated protein kinase pathway and regulates actin cytoskeletal organization"		1-10,17
	JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 275, no. 6, 11 February 2000 (2000-02-11), pages 4311-4322, XP002285646 ISSN: 0021-9258 cited in the application the whole document		
X	DATABASE GENESEQ 'Online! 8 February 2001 (2001-02-08) SHIMKETS RA, LEACH M: "Human ORFX ORF1427 polypeptide sequence SEQ ID NO:2854" Database accession no. AAB41663 XP002285651 the whole document		1-10,17
<b>X</b>	DATABASE EMBL 'Online! 9 February 1999 (1999-02-09) NAGASE T. ET AL.,: "Homo sapiens mRNA for KIAA0881 protein, partial cds." Database accession no. AB020688 XP002285652 cited in the application		1-10,17
X	the whole document  DATABASE EMBL 'Online! 15 October 2001 (2001-10-15) JENKINS SG, ET AL., : "Homo sapiens serine/threonine kinase TAO1 mRNA, complete cds." Database accession no. AY049015 XP002285653 the whole document		1-10,17
X	DATABASE EMBL 'Online! 14 March 2000 (2000-03-14) NAGASE T. ET AL., : " Homo sapiens mRNA for KIAA1361 protein, partial cds." Database accession no. AB037782 XP002285654 cited in the application abstract		1-10,17
A	US 6 034 228 A (MOORE WILLIAM CRAIG ET AL) 7 March 2000 (2000-03-07) the whole document		



		FET/EP 03	14674
C.(Continua	etion) DOCUMENTS CONSIDERED TO BE RELEVANT	THE STATE OF THE S	
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
A	HIROSUMI JIRO ET AL: "A central role for JNK in obesity and insulin resistance." NATURE (LONDON), vol. 420, no. 6913, 21 November 2002 (2002-11-21), pages 333-336, XP002285647 ISSN: 0028-0836 (ISSN print) cited in the application the whole document		
A	OGG S ET AL: "THE FORK HEAD TRANSCRIPTION FACTOR DAF-16 TRANSDUCES INSULIN-LIKE METABOLIC AND LONGEVITY SIGNALS IN C. ELEGANS"		
	NATURE, MACMILLAN JOURNALS LTD. LONDON, GB, vol. 389, 30 October 1997 (1997-10-30),		
	pages 994-999, XP002926981 ISSN: 0028-0836 cited in the application the whole document		
	WO 00/01846 A (MORTIER KATHERINE ;DEVGEN NV (BE); BOGAERT THIERRY (BE); PLAETINCK) 13 January 2000 (2000-01-13) cited in the application		
<b>Р,</b> Х	WO 03/051905 A (EXELIXIS INC) 26 June 2003 (2003-06-26) see human TAOJIK DNA #s 1, 4, 8 the whole document		1-10,17
		٠., ٠	
i			*
	-		
			and the second
-			



Box I Observations where certain claims were found unsearchable (Continua	ation of item 1 of first sheet)
en de la companya de La companya de la comp	
This International Search Report has not been established in respect of certain claims under A	rticle 17(2)(a) for the following reasons:
	The state of the s
Claims Nos.:      because they relate to subject matter not required to be searched by this Authority, na	amely:
	A Company of the Carlo
2. X Claims Nos.: 11-16	en e
because they relate to parts of the International Application that do not comply with the an extent that no meaningful International Search can be carried out, specifically:	e prescribed requirements to such
see FURTHER INFORMATION sheet PCT/ISA/210	
	gradient was a state of the second
Claims Nos.:  because they are dependent claims and are not drafted in accordance with the second	d and third sentences of Rule 6.4/a)
Box II Observations where unity of invention is lacking (Continuation of item	2 of Erst shoot)
box ii Observations where unity of invertibilitis facking (Continuation of item	2 Of Hist Sheet)
This International Searching Authority found multiple inventions in this international application,	as follows:
see additional sheet	
out add votolial offices	
and the second of the second o	
por la completation de la completa d La completa de la co	
As all required additional search fees were timely paid by the applicant, this Internation searchable claims.	nal Search Report covers all
2. X As all searchable claims could be searched without effort justifying an additional fee, t	his Authority did not invite nevment
of any additional fee.	ins Additing did not hivre payment
As only some of the required additional search fees were timely paid by the applicant, covers only those claims for which fees were paid, specifically claims Nos.:	this International Search Report
details any those orange for which lees were paid, specifically claims (vos	
4. No required additional search tees were timely paid by the applicant. Consequently, It	nis International Search Report is
4. No required additional search fees were timely paid by the applicant. Consequently, the restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	no memanona ocusin risportis
	•
Remark on Protest The additional search fees were a	accompanied by the applicant's protest.
No protest accompanied the payr	nent of additional search fees.

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box 1.2

Claims Nos.: 11-16

Present claims 11-16 relate to a compound defined by reference to a desirable characteristic or property, namely to be a modulator of a protein or polypeptide according to claim 8.

The claims cover all compounds having this characteristic or property, whereas the application provides support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for none of such compounds. In the present case, the claims so lack support, and the application so lacks disclosure, that a meaningful search over the whole of the claimed scope is impossible. Independent of the above reasoning, the claims also lack clarity (Article 6 PCT). An attempt is made to define the compounds by reference to a result to be achieved. Again, this lack of clarity in the present case is such as to render a meaningful search over the whole of the claimed scope impossible.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-17 (all partially)

Nucleic acid encoding the amino acid sequence of SEQ ID NO: 2, and subject-matter related thereto.

2. Claims: 1-17 (all partially)

Nucleic acid encoding the amino acid sequence of SEQ ID NO: 4, and subject-matter related thereto.

3. Claims: 1-17 (all partially)

Nucleic acid encoding the amino acid sequence of SEQ ID NO: 6, and subject-matter related thereto.

4. Claims: 1-17 (all partially)

Nucleic acid encoding the amino acid sequence of SEQ ID NO: 8, and subject-matter related thereto.

5. Claims: 1-17 (all partially)

Nucleic acid encoding the amino acid sequence of SEQ ID NO: 10, and subject-matter related thereto.

International Application No. PEP/EP 03/14674

	_,				<del></del>
	Patent document cited in search report	Publication date	Patent family member(s)		Publication date
	WO 9953036	A 21-10-1999	AU 364249	9 A	01-11-1999
	ON UK FILE		CA 236917	2 A1	21-10-1999
:	9,2		EP 107372	3 A2	07-02-2001
1			JP 200252200	9 T	23-07-2002
ŧ			WO 995303	6 A2	21-10-1999
			US 200305023	0 A1	13-03-2003
		-	US 665671	6 B1	02-12-2003
	US 6034228	A 07-03-2000	US 630009	8 B1	09-10-2001
	WO 0001846	A 13-01-2000	AU 76922	3 B2	22-01-2004
Ì		Å.	AU 490799	9 A	24-01-2000
		•	BR 991180	2 A	22-01-2002
			CA 233261	9 A1	13-01-2000
	•		CN 132335		21-11-2001
			DE 2992429	8 U1	12-09-2002
			DE 2992429		12-09-2002
1	. •		DE 109352	6 T1	11-10-2001
l ·			WO 000184	6 A2	13-01-2000
			EP 119756		17-04-2002
1			EP 109352	6 A2	25-04-2001
<u> </u>			GB 234988		15-11-2000
			GB 236288	5 A ,B	05-12-2001
ļ		,	GB 237027	5 A ,B	26-06-2002
		•	HK 102914	2 A1	25-04-2003
}			HU 010357	1 A2	28-01-2002
l	•		JP 200251907	2 T	02-07-2002
			NO 2001001		05-03-2001
1 .			NZ 50918	2 A	30-01-2004
l			PL 34797	8 A1	06-05-2002
			US 200306162	6 A1	27-03-2003
			ZA 20000765	3 A	19-09-2002
	WO 03051905	A 26-06-2003	CA 246080		26-06-2003
			WO 0305190		26-06-2003
			US 200315753	1 A1	21-08-2003
ŀ					







Application No: Claims searched:

GB 0230014.3

1-10 & 17 (in part)

Examiner: Date of search:

Dr Rowena Dinham

18 June 2003

## Patents Act 1977: Search Report under Section 17

#### Documents considered to be relevant:

Category	Relevant to claims	Identity of document and passage or figure of particular relevance		
Х	1-10 & 17	Gene; Vol 29, pp 137-147 (2001). Berman et al. "kin-18, a C-elegans protein kinase involved in feeding". See entire document, especially Figure 1		
X	1-10 & 17	WO 99/53036 A2 (SUGEN, INC.) See especially page 46 line 15- page 47 line 20 and SEQ ID No 90		
Х	1-10 & 17	J Biol Chem; Vol 273, pp 28625-28632 (1998). Hutchison et al.  "Isolation of TAO1, a protein kinase" See entire document, especially Figure 1B		
Х	1-10 5 & 17	J Biol Chem; Vol 274, pp 33287-33295 (1999). Tassi et al. "Human JIK, a novel member of the STE20 kinase family" See entire document, especially Figure 1B		
Х	1-10 ; & 17	GenBank Accession number U32275. See entire document		

#### Categories:

x	Document indicating lack of novelty or inventive step	Α	Document indicating technological background and/or state of the art.
Y	Document indicating lack of inventive step if combined with one or more other documents of same category.	P	Document published on or after the declared priority date but before the filing date of this invention.
&	Member of the same patent family	Е	Patent document published on or after, but with priority date earlier than, the filing date of this application.

#### Field of Search:

Search of GB, EP, WO & US patent documents classified in the following areas of the UKC<sup>v</sup>:

Worldwide search of patent documents classified in the following areas of the IPC<sup>7</sup>:

C07K; C12N

The following online and other databases have been used in the preparation of this search report:

WPI, JAPIO, EPODOC, BLASTn, BLASTp, MEDLINE, BIOSIS